## In the Claims

- 1. (Withdrawn) DNA comprising one or more genes specific for 5S clavam biosynthesis in *S. clavuligerus* and which is not essential for 5R clavam biosynthesis.
- 2. (Withdrawn) DNA according to claim 1 as identified in Figure 1 (SEQ ID No: 1).
- 3. (Withdrawn) DNA according to claim 1 having the sequence or substantially the sequence shown in Figure 1 as orfup3, orfup2, orfup1, orfdwn1, orfdwn2 or orfdwn3 (SEQ ID Nos: 2-7).
- 4. (Withdrawn) DNA according to claim 1 having the sequence or substantially the sequence shown in Figure 1 as orfup1 (SEQ ID No: 4).
- 5. (Withdrawn) DNA which hybridises under conditions of high stringency with the DNA of claim 1.
- 6. (Withdrawn) A vector comprising the DNA of claim 1 in which one or more of the genes specific for 5S clavam biosynthesis has been disrupted or otherwise made defective.
- 7. (Withdrawn) A vector according to claim 6 containing one or more defective genes which is pCEC060, pCEC061, pCEC056 or pCEC057.
- 8. (Withdrawn) A vector according to claim 7 which is pCEC061.
- 9. (Withdrawn) A host containing the vector of claim 6.
- 10. (Withdrawn) A host according to claim 9 which is capable of producing raised levels of clavulanic acid.
- 11. (Withdrawn) A host according to claim 9 which is capable of producing low or no levels of 5S clavam.
- 12. (Withdrawn) A host according to claim 9 which is S. clavuligerus.

13. (Withdrawn) S. clavuligerus comprising DNA corresponding to an open reading frame flanking cas1 which DNA has been disrupted or otherwise made defective.

- 14. (Withdrawn) *S. clavuligerus* according to claim 13 wherein the open reading frame is selected from the group consisting of orfup3, orfup2, orfup1, orfdwn1, orfdwn2 and orfdwn3.
- 15. (Withdrawn) A process for improving 5R clavam production in a suitable microorganism comprising manipulation of DNA as defined in claim 1 and its inclusion in the microorganism.
- 16. (Withdrawn) A process according to claim 15 wherein said suitable microorganism is *S. clavuligerus*.
- 17. (Withdrawn) A process for improving 5R clavam production in *S. clavuligerus* comprising disrupting or otherwise making defective DNA regions flanking cas1.
- 18. (Withdrawn) A process according to claim 15 wherein said DNA corresponds to open reading frames selected from the group consisting of orfup3, orfup2, orfup1, orfdwn 1, orfdwn2 and orfdwn3.
- 19. (Withdrawn) A process according to claim 15 wherein said DNA corresponds to open reading frame orfup1.
- 20. (Withdrawn) A process according to claim 15 wherein said 5R clavam is clavulanic acid.
- 21. (Withdrawn) A process for the identification of a microorganism suitable for high 5R clavam production comprising a preliminary screening for microorganisms with low or no 5S clavam production.

- 22. (Withdrawn) A process according to claim 21 wherein the microorganism is S. clavuligerus.
- 23. (Withdrawn) A process according to claim 22 wherein the 5R clavam is clavulanic acid.
- 24. (Withdrawn) A process according to claim 21 wherein one or more genes specific for the production of 5S clavams is defective.
- 25. (Withdrawn) A microorganism which is capable of 5R clavam production and low or no 5S clavam production obtainable by the process of claim 15.
- 26. (Withdrawn) A microorganism obtainable by the process of claim 25 which is capable of producing clavulanic acid but which does not produce clavam-2- carboxylate.
- 27. (Withdrawn) A microorganism obtainable by the process of claim 25 which is capable of producing clavulanic acid but which does not produce 2-hydroxymethylclavam.
- 28. (Withdrawn) A microorganism obtainable by the process of claim 25 which is capable of producing clavulanic acid but which does not produce clavam-2- carboxylate and 2-hydroxymethylclavam.
- 29. (Withdrawn) A microorganism obtained by the process of claim 15 which is strain 56-1A, 56-3A, 57-2B, 57-1C, 60-1A, 60-2A, 60-3A, 61-1A, 61-2A, 61-3A or 61-4A.

Claims 30-36 (Cancelled).

37. (Withdrawn) A process for the preparation of a composition comprising potassium clavulanate and amoxycillin which process comprises producing clavulanic acid from a microorganism according to claim 25 and thereafter converting it to the potassium salt and combining the potassium salt with amoxycillin.

45. (Presently Amended) A pharmaceutical composition which is free of one or more of the clavams selected from the group consisting of clavam-2-carboxylate, 2-hydroxymethylclavam, and 2-(3-alenylalanyl)clavam comprising clavulanic acid, obtainable by the process comprising the steps of:

- (a) disrupting or otherwise making defective a DNA comprising one or more genes specific for 5S clavam biosynthesis in <u>S. clavuligerus</u> a <u>Streptomyces</u>, selected from the group consisting of orfup3 (SEQ ID NO: 2), orfup2 (SEQ ID NO: 3), orfup1 (SEQ ID NO: 4), orfdwn1 (SEQ ID NO: 5), orfdwn2 (SEQ ID NO: 6) and or orfdwn3 (SEQ ID NO: 7) or a degenerate variant of orfup3 (SEQ ID NO: 2), orfup2 (SEQ ID NO: 3), orfup1 (SEQ ID NO: 4), orfdwn1 (SEQ ID NO: 5), orfdwn2 (SEQ ID NO: 6) or orfdwn3 (SEQ ID NO: 7);
  - (b) fermenting said <u>S. clavuligerus</u> a <u>Streptomyces</u>; and
  - (c) purifying clavulanic acid therefrom.
- 46. (Cancelled)
- 47. (Presently Amended) The pharmaceutical composition according to claim <u>44 45</u>, wherein the clavulanic acid is in the form of a potassium salt.
- 48. (Presently Amended) The pharmaceutical composition according to claim <u>44 45</u>, further comprising a beta-lactam antibiotic.
- 49. (Presently Amended) The pharmaceutical composition according to claim 4748, wherein the beta-lactam antibiotic is amoxycillin.
- 50. (New) *Streptomyces* culture media comprising clavulanic acid and free of any 5S clavam as measured by HPLC analysis.
- 51. (New) The *Streptomyces* culture media according to claim 50 wherein the 5S clavam is selected from the group consisting of: clavam-2-carboxylate (C2C), 2-hydroxymethylclavam (2HMC), and 2-(3-alanyl)clavam (AC).